



Stereotactic radiotherapy for the treatment of pulmonary metastases: a narrative review

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Objective: To describe the rationale and the technique summarizing the most recent and relevant results reported in the scientific literature on stereotactic radiotherapy for pulmonary metastases.

Background: Stereotactic body radiation therapy (SBRT) is a radiation therapy technique that delivers a high dose of radiation very precisely to an extracranial target, using either a single dose or a small number of fractions. The lung is the main site of metastatic spread for most solid tumors and local treatments have an emerging role, especially in oligometastatic patients. SBRT requires a coordinated team effort between radiation oncologists, medical physicists, radiation therapists and nurses. With this review, we present and analyze the most recent data in terms of clinical results and toxicity that also in the absence of phase III randomized trial allow to reach clinically relevant findings on the cost-benefit balance of stereotactic radiotherapy of lung metastasis (LM).

Methods: In this review, we analyzed the literature data of the last ten years on patients treated with SBRT for lung oligometastasis. We found a number of non-randomized retrospective studies heterogeneous in terms of primary tumour, number of lesions and techniques.

Conclusions: All authors conclude that SBRT represents an effective, safe option, with a low toxicity profile, which improves local control of the disease and the overall survival (OS) of patients with lung metastases. These characteristics together with the convenience of the few treatment sessions make it an excellent alternative to other more invasive therapy. Future randomized studies will enable people to better define the most appropriate dose and fractionation, the most suitable patients in terms of histology of the primary tumors and the best integration with systemic therapies.

Keywords: Radiotherapy; stereotactic radiotherapy; pulmonary metastasis; oligometastasis

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Introduction

The lung is the main site of metastatic spread for most solid tumors and local treatments have an emerging role especially in oligometastatic patients. Surgical resection has

been widely used in carefully selected patients (1).

Stereotactic body radiation therapy (SBRT) is an external beam radiation therapy technique that very precisely deliver a high dose of radiation to an extracranial target within

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the body, using either a single dose or a small number of fractions. Specialized treatment planning results in steep dose gradients beyond the target (2).

The European Organization for Research and Treatment of Cancer (EORTC) Lung Cancer Group agrees that the definition of oligometastatic disease (OMD) is appropriate when a radical treatment is technically feasible. The potential toxicity should be acceptable, with all sites being amenable to a local treatment that may modify the course of the disease and being an opportunity for long-term disease control.

Extensive diagnostic evaluation including positron emission tomography/computed tomography (PET/CT) and brain imaging, are necessary to define a patient as oligometastatic (3). A maximum of five metastases and three organs has been proposed (not including mediastinal lymph nodes). The presence of diffuse serosal metastases (meningeal, pericardial, pleural, and mesenteric) or bone marrow involvement are excluded from this definition.

In a recently ESTRO-ASTRO published consensus document has been given a new definition for OMD that is independent of the type and histology of primary tumor and of the site of metastases but includes only 1–5 treatable lesions with or without a controlled primary site of disease (4). Determinants, however, are the number of lesions (regardless of the localization) and the radical purpose of the therapeutic intent.

Diagnostic imaging should be performed using any modalities adequate to scan and identify any metastases detecting also small lesions. The most common are computed tomography (CT) scan of the chest/abdomen/pelvis, magnetic resonance imaging (MRI) of the brain or spine and whole body PET/CT. SBRT allowing precise delivery of high radiation dose, defined as “ablative dose”, with maximum sparing of normal tissue has emerged as a safe and efficacious treatment modality for selected patients with pulmonary metastases (PM) (5,6).

In order to consider the dose effective and therefore “radical” in SBRT, using the concept of BED (biologically effective dose), a minimum BED ($\alpha/\beta = 10$) of 100 gray (Gy) is conventionally considered necessary in order to achieve local control of OMD. BED is the expression of the combination of dose per fraction, total dose and tissue characteristics (α/β ratio of the specific tissue). Therefore is a measure of the true biological dose delivered and is commonly used to compute and compare the iso-effective dose (7). α/β ratio represent the intrinsic radio sensitivity of the irradiated tissue: most tumors exhibit an high α/β ratios (e.g., $\alpha/\beta = 10$ Gy is commonly used) (8).

The radiobiological advantage of SBRT depend on the shorter treatment time and the high dose per fraction compared to conventional radiotherapy, preventing the cellular repopulation that can occur during a longer radiation therapy course (9).

The best approach to cancer patients is multidisciplinary with the involvement of the different professional figures such as surgeons, medical oncologists, radiation oncologists, radiologists, pathologists and specialist in nuclear medicine. In specific situation, other professional figures are involved, such as psychologists, nutritionists and psychiatrists, in order to adequately identify the individual clinical scenario and offer the most appropriate treatment, taking into account patient and disease characteristics, clinical history and previous treatments. SBRT requires a coordinated team effort between radiation oncologists, medical physicist, radiation therapists and nurses (5).

The procedure starts with the CT simulation of the immobilized patient in order to identify the target volume and the nearby normal organs that should receive the minimum dose, named organ at risk (OAR), and to study the respiratory movement. On the CT simulation images, with the aid of dedicated software, the radiation oncologist defines the target volume and OAR and prescribes the radiation dose. The medical physicist prepares the treatment plan and refine it with the radiation oncologist that ultimately give his approval. The first day of the treatment, all that has been previously planned is verified with the patient at the linear accelerator and the treatment is delivered. The essential point of the process is the precise daily reproducibility of the treatment. This is achieved with a compromise between a tight fitting and a comfortable immobilization of the patient often achieved with the creation of a customized thermoplastic masks, but also by the motion management and by the use of image-guided verification systems.

All these measures enable to reduce intrafraction/interfraction variability, therefore the safety margins conventionally given to the target volume could be decreased. This enables to reduce the irradiated volume of OAR ultimately allowing the increase of the prescribed dose.

A PET-FDG simulation can be associated to the CT-simulation with the same immobilization system taking advantage of metabolic images in order to obtain a better definition of the target. Moreover, inoculating contrast media to the patient during the CT simulation, which is normally performed without it, help in selected cases as perimediastinal lesions, a finest identification of the target.



Figure 1 Patient immobilized with thermoplastic mask and diaphragmatic frame for abdominal compression built according to the patient's rib arch by the radiation therapists. The figure shows the respiratory gating system adopted in our department. Gating is a system that tracks a patient's normal respiratory cycle with a chest/abdomen marker camera. This image is published with the patient's consent.

In order to clear up the procedure described above; we report here a graphic depiction of the sequence of event that led to stereotaxic radiotherapy treatment performed at the Radiotherapy Department of "Istituto Clinici Scientifici Maugeri IRCCS" of Pavia. Initially the anatomic region of interest is scanned with the CT simulator while the motion of the markers placed on chest and abdomen is tracked with an infrared camera (*Figure 1*). The images are analyzed using four dimensions (4D) software (the fourth dimension is time) to accurately correlate the shift of the tumor with the breathing motions. The target volume will be defined by the envelope of all different position of the lesion during breathing motion (*Figure 2*).

With this review, we present and analyze the most recent data in terms of clinical results and toxicity published in the literature on the topic of stereotactic radiotherapy of lung metastasis (LM). We present the following article in accordance with the Narrative Review reporting checklist (available at <https://asj.amegroups.com/article/view/10.21037/asj-21-53/rc>).

Methods

In order to collect and analyze the most significant papers

of the last ten years starting from 2011 forward, we carried out a literature search using the terms "stereotactic body radiotherapy" and "pulmonary oligo metastases".

We selected 31 articles considering the following points: originality of the study, appropriate prescribed doses excluding those with only palliative intent, employed fractionation (1 to 10 fractions) and clinical outcomes.

The analysis showed two differences in the treatment approach in terms of the number of fractions and technique: single (*Table 1*) *vs.* multiple fractions (*Table 2*) and conformal three-dimension (3D) techniques *vs.* volumetric modulated arc therapy (VMAT).

Discussion

The pattern of care of PM with stereotaxic radiotherapy vary in doses and number of fractions. We begin by discussing the experiences in single fraction moving then on those in multiple fractions.

We analyzed the literature of the last 10 years, starting in 2011. All studies are retrospective analysis of patients treated in the early years with conformal 3D radiotherapy and more recently with VMAT.

The group of Ricardi *et al.* (6) gave an important

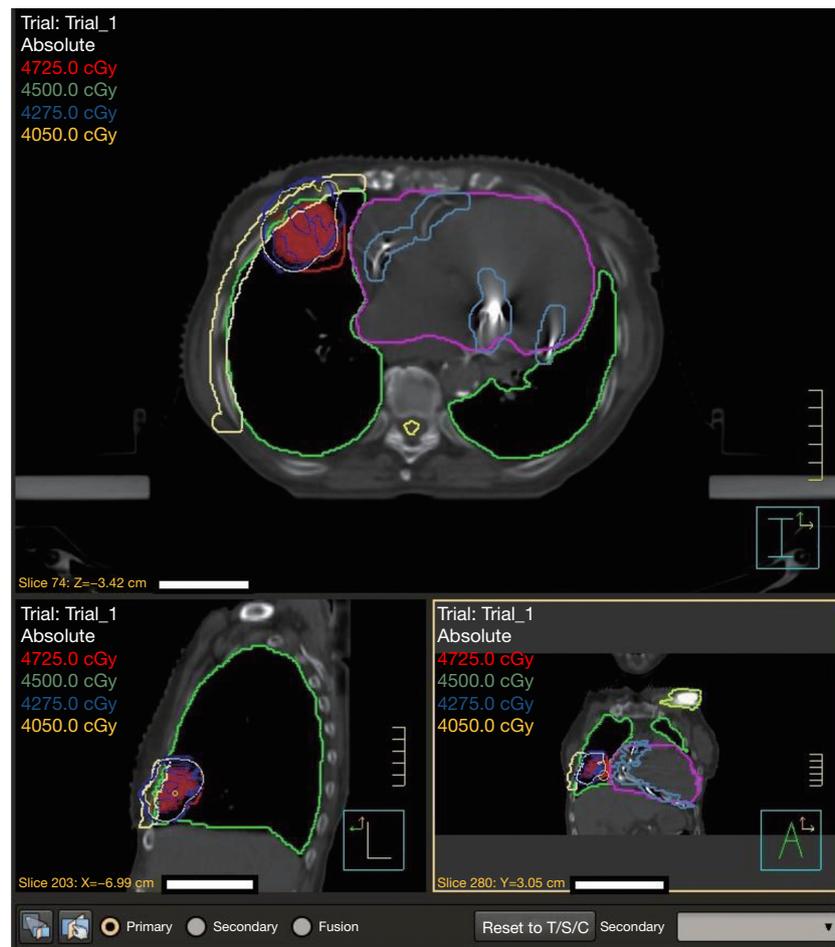


Figure 2 Stereotactic treatment plan with isodose distribution on a lesion of the right lobe on coronal, sagittal and axial plane of the CT simulation.

contribution in this field publishing in 2012 a study on 77 metastases, 51 of which treated with 26 Gy single fraction and 26 with 45 Gy in 3 fractions. Results at two years were excellent in terms of local control (LC), and overall survival (OS) in a context of very low rate of grade 3 (G3) toxicity. These results were confirmed with an update published two years later by Filippi *et al.* (11) that reported the outcome of treatment of 90 metastases on 67 patients. The primary tumors were various but the vast majority were colorectal cancer. The prescribed dose was 26 Gy in single fraction. Two-year LC and OS were comparable to the older study. This study is appealing for the uniformity of the dose prescribed, on a relevant number of patients with an adequate follow up (FU) for a preliminary estimate of LC and OS.

The same author in 2016 (15) published data on 43 lesions mostly treated with single fraction as shown in *Table 1*.

LC and OS were good, 86% and 77% respectively with no G3 toxicity. The aim of the study was also to compare the outcome of stereotactic radiotherapy with surgery. Even with all the limits of the retrospective observational analysis and the difference in the samples size, 142 Surgery *vs.* 43 SBRT, the author concludes that OS at 2 years are comparable in the two groups.

Osti *et al.* in 2013 (10) conducted a study on 66 patients, 103 metastases from different primary (most were from lung, breast and rectal cancers). At 1 and 2 years LC were 89.1% and 82.1%, OS 76.4% and 31.2%, cancer-specific survival (CSS) 78.5% and 35.4%, progression-free survival (PFS) 53.9% and 22%, respectively. Toxicity profiles was good, with two Grade 3 toxicity (pneumonitis). The same author in 2018 (16) published the data on 166 lesions, mostly from lung and colorectal primary tumor, treated

Table 1 Characteristics of the selected studies in single fraction studies.

Author, year	No. of patients	No. of lung lesions	Primary site	Dose	Local control	Overall survival	Toxicity, G ≥3
Ricardi <i>et al.</i> (6), 2012	61	77	All (mostly lung)	26 Gy/1 fr (51/77); 45 Gy/3 fr (26/77)	2-y 89%	2-y 67%	2% G3 pneumonitis
Osti <i>et al.</i> (10), 2013	66	103	All (mostly lung, rectal and breast)	23 Gy 1/fr (central); 30 Gy 1/fr (peripheral)	2-y 82%	2-y 31%	3% G3 pneumonitis
Filippi <i>et al.</i> (11), 2014	67	90	All (mostly lung and colorectal)	26 Gy 1 fr	2-y 88%	2-y 71%	Not reported
Wang <i>et al.</i> (12), 2015	95	134	All (mostly lung)	30–60 Gy/1–5 fr	2-y 91%	2-y 61%	3% G3 pneumonitis
Siva <i>et al.</i> (13), 2015	65	85	All	18–26 Gy/1 fr (41/85); 48–50 Gy/4–7 fr (44/85)	2-y 93%	2-y: 71%	None
Davis <i>et al.</i> (14), 2015	64	66	All (mostly colon)	16–60 Gy/1–5 fr	2-y 70%	2-y 50%	None
Filippi <i>et al.</i> (15), 2016	28	43	Colorectal	26 Gy/1 fr (31/43); 25 Gy/3 fr (8/43); 55–60 Gy/8–10 fr (4/43)	Median FU 27-m: local recurrence 21%	2-y 77%	None
Osti <i>et al.</i> (16), 2018	129	166	All (mostly lung and colorectal)	30 Gy/1 fr	LPFS: 3-y 80.1%; 5-y 79.2%	3-y 50.7%; 5-y 37.9%	3.6% G3 pneumonitis; 1.2% G5 pneumonitis; 7.4% G3 lung fibrosis; 1.3% rib fracture
Kalinauskaite <i>et al.</i> (17), 2020	52	94	All (mostly colorectal)	22–8 Gy/1–5 fr (mostly 24 Gy/1 fr)	1-y and 2-y for SFRS vs. fSBRT: 89% and 83% vs. 75% and 59%, respectively	1-y 84%; 2-y 71%	None

fr, frequency; y, year; FU, follow up; m, month; LPFS, local relapse-free survival; SFRS, single fraction radiosurgery; fSBRT, fractionated stereotactic body radiotherapy.

with 30 Gy single fraction. The 3 and 5 years local relapse-free survival (LPFS) were 80.1% and 79.2% (median not reached), respectively. One hundred and forty-eight patients were evaluated for late toxicity, see *Table 1*. The 3- and 5-years OS were 50.7% and 37.9%, respectively. The data of these two studies are interesting for the homogeneity of the dose delivered and certainly constitute a home base for future study on the radiosurgical approach to pulmonary oligometastases.

Wang *et al.* (12) in 2015 published the results of stereotactic radiotherapy of 134 lung lesions of varying primary tumors, the majority lung cancer. The doses were 30–60 Gy in 1–5 fractions. G3 toxicities were low (3% of G3 pneumonitis). At two years, LC and OS were 91% and 61%, respectively.

Also in 2015, Siva *et al.* (13) published a study on 85 LM of varying primary tumors. They reported a variety

of approaches that included single fraction for 41 of the 85 lesions with a dose between 18 and 26 Gy, and 4 to 7 fractions for 44 of the 85 lesions, dose range between 45 and the 50 Gy. They reported no relevant toxicities with a two-year LC of 93% and an OS of 72%.

More recently, in 2020, Kalinauskaite *et al.* (17) searched for prognostic features of better survival outcomes after single fraction radiosurgery (SFRS) and fractionated SBRT (fSBRT) in patients with lung oligometastases. They treated fifty-two patients with SFRS or fSBRT. Median dose to the planning target volume (PTV) for SFRS was 24 Gy (range, 17–26 Gy) compared to 45 Gy (range, 20–60 Gy) in 2–12 fractions for fSBRT. LC and OS at 1 and 2 years were better for SFRS *vs.* fSBRT. LM treated with SFRS were significantly smaller (P=0.001). Longer time to first metastasis, good KPS (<70) and N0 predicted better OS.

Our review aim to describe the role of SBRT even in

Table 2 Characteristics of the selected studies in multiple fraction studies

Author, year	No. of patients	No. of lung lesions	Primary site	Dose	Local control	Overall survival	Toxicity, G \geq 3
Takeda <i>et al.</i> (18), 2011	34	44	All (mostly colon cancer)	50 Gy/5 fr	2-y 72% (colon) 94% (other)	Not reported	2% G3 pneumonitis
Oh <i>et al.</i> (19), 2012	57	67	All (mostly lung)	50–60 Gy/5–4 fr	3-y 94.5%	2-y 60%	1 G5 pneumonitis
Inoue <i>et al.</i> (20), 2013	87	189	All (mostly colorectal)	48 Gy/4 fr (87/189); 52–60 Gy/10 fr (36/189); Other (66/189)	2-y 80%	2-y 47%	9% G3, 1% G4 pneumonitis
Baschnagel <i>et al.</i> (21), 2013	32	47	All	Various, mostly 60 Gy/5 fr (36/47)	2-y 92%	2-y 76%	16% G3 (various)
Widder <i>et al.</i> (22), 2013	42	Not reported	All (mostly colorectal)	60 Gy/3 fr (23/42); 60 Gy/5 fr (9/42); 60 Gy/8 fr (10/42)	2-y 84%	2-y 86%	Not reported
Navarria <i>et al.</i> (5), 2014	76	118	All (mostly colorectal)	60 Gy/3 fr (peripheral); 48 Gy/4 fr (peripheral >2 cm); 60 Gy/8 fr (central)	2-y 89%	2-y 73%	None
Soyfer <i>et al.</i> (23), 2014	22	53	Sarcoma	Various, mostly 60 Gy/3 fr (33/53)	Mean 5.1-m: 64% (0–10 mm); 30% (11–20 mm); 6% (>20 mm)	5-y 62%	5% pneumonitis
Garcia-Cabezas <i>et al.</i> (24), 2015	42	53	All (mostly colon)	60 Gy/5 fr (35/53); 60 Gy/8 fr (11/53); 50 Gy/10 fr (7/53)	2-y 87%	2-y 60%	None
Qiu <i>et al.</i> (25), 2015	65	Not reported	All	50 Gy/5–10 fr	2-y 31%	2-y 43%	Not reported
Jung <i>et al.</i> (26), 2015	50	79	Colon	40–48 Gy/4 fr (56/79); 60 Gy/3–4 fr (23/79 fr)	3-y 71%	3-y 64%	None
Aoki <i>et al.</i> (27), 2016	66	76	All (mostly lung)	45–60 Gy/5–9 fr	3-y 91%	3-y 76%	None
Guckenberger <i>et al.</i> (28), 2016	397	525	All	Various	Median FU 16-m: 86.7%	Not reported	Not reported
Lischalk <i>et al.</i> (29), 2016	20	Not reported	All (mostly lung)	35–40 Gy/5 fr	2-y 57%	2-y 40%	5% G3 pneumonitis 5% G4 atelectasia
Baumann <i>et al.</i> (30), 2016	30	39	Sarcoma	50 Gy/4–5 fr (32/39); <50 Gy (7/39)	2-y 86%	2-y 43%	None
Rieber <i>et al.</i> (31), 2016	700	Not reported	All	Median number of fractions: 3; Median single doses: 12.5 Gy (range 3.0–33.0 Gy)	1-y 90.9%; 2-y 81.2%	1-y 75.1%; 2-y 54.4%	6.5% \geq G2 pneumonitis; 0.2% G5 pneumonitis; 3.2% other \geq G2+ toxicities
Ricco <i>et al.</i> (32), 2017	304	327	All	Median 50 Gy /3–5 fr	1-y 80.4%; 3-y 58.9%; 5-y 46.2%	1-y 74.1%; 3-y 33.3%; 5-y 21.8%	Not reported
Franceschini <i>et al.</i> (33), 2017	200	Not reported	All (mostly colorectal)	30–60 Gy/1–8 fr	1-y 91%; 2-y 84.9%	1-y 88.7%; 2-y 65.4%	None

Table 2 (continued)

Table 2 (continued)

Author	No. of patients	No. of lung lesions	Primary site	Dose	Local control	Overall Survival	Toxicity, G ≥3
Mazzola <i>et al.</i> (34), 2018	78	114	All (mostly lung or colon tumor)	SBRT with a biologically equivalent dose (BED) ≥100 Gy	Median local failure free survival: 18 m	Median 19.6 m	None
Lindsay <i>et al.</i> (35), 2018	44	117	Sarcoma	36–50 Gy/6–10 fr (mostly 50 Gy/10 fr)	2-y 95%	2-y 82%; 5-y 50%	1 Esophageal stricture
Hörner-Rieber <i>et al.</i> (36), 2019	301	336	NSCLC	Various	1-y 91.9%; 2-y 82.0%; 3-y 76.4%; 5-y 70.3%	1-y 80.8%; 2-y 62.2%; 3-y 48.1%; 5-y 28.8%	Not reported
Wegner <i>et al.</i> (37), 2019	44	50	All (mostly colorectal)	48 Gy/5 fr	1-y 82%	1-y 66%	None
Berkovic <i>et al.</i> (38), 2020	104	132	All (mostly lung, gastrointestinal)	Various, mostly 60 Gy/3 fr (73 lesions)	1-y 89.3%; 2-y 80%; 3-y 77.8%	1-y 92.2%; 2-y 80.9%; 3-y 72.0%	3 G3 pneumonitis and 1 G4
Kessel <i>et al.</i> (39), 2020	219	316	All	Various median dose 35 Gy and median dose per fraction 8 Gy	1-y 92%; 2-y 84%; 3-y 78%; 4-y 77%; 5-y 75%; 10-y 75%	1-y 74%; 2-y 54%; 3-y 39%; 4-y 29%; 5-y 26%; 10-y 17%	G3 2.9%

fr, frequency; y, year; FU, follow up; m, month; SBRT, stereotactic body radiotherapy; NSCLC, non-small cell lung cancer.

studies in which the treatment was fractionated. Also in these studies the results were satisfactory and the toxicity low. In 2011, Takeda *et al.* (18) published the results of the treatment of 44 secondary lesions of various primary tumors (mostly colon cancers) after 50 Gy in 5 fractions. Remarkable for this analysis is the difference of LC based on the primary tumor: when colon cancer was the primary, the two-year LC was much lower compared to other primary.

Oh *et al.* (19) described similar results in his analysis on 67 lesions with lung cancer as predominant primary site. Three-year LC rate was 94.5%, two-year OS 60%, reported toxicity was 13% for rib fracture and 1 death from pneumonitis.

In 2013, Inoue *et al.* (20) published data on a larger sample of patients treated for oligo metastases mostly from colorectal cancer. They treated 189 lesions with various fractionation schemes (see Table 2); the most represented in the analysis was 48 Gy/4 fractions. The two-year LC was 80%, OS 47% and a low toxicity profile.

One preliminary observation that could be drawn from the studies reported above, though conducted on different

samples and with different fractionation and doses, is that treating metastases from colorectal cancer achieve lower LC compared to other primary tumors.

In 2014, Navarria *et al.* (5) published data of treatment outcome of 118 metastases (mostly from colorectal cancer). The total dose prescribed varied according to tumor location and size: 48 Gy in 4 fractions for peripheral lesions, the schedule adopted most frequently, 60 Gy in 8 fractions for central lesions and 3 fractions for peripheral lesions with diameter ≤2 cm. LC at 1, 2 and 3 years was 95%, 89% and 89% respectively. OS at 1, 2 and 3 years was 84.1%, 73% and 73% respectively. No major pulmonary toxicity, chest pain or rib fracture occurred.

Three years later the same Institution published the results of further experiences. Franceschini *et al.* (33) described the stereotactic treatment of 200 patients with a predominance of colorectal cancer (49.5% of the cases) as primary tumor. This experience with a large number of patients with metastases from colorectal carcinoma confirmed his “radio-resistant” trait negatively affecting LC.

Also, the analysis of AoKi *et al.* (27) support this hypothesis. They report of 76 oligo-recurrences in the

lungs treated with SBRT from various primary sites. The median SBRT dose was 50 Gy (range, 45–60 Gy) given in a median of five (range, 5–9) fractions. Oligo-recurrence from colorectal cancer had worse outcomes for LC and DFS compared with those of other primary cancer sites.

In order to obtain more reliable information to guide the choice of the most appropriate therapy, is of great value the availability of study not only with well selected patient but also highly homogeneous in term of histology of the primary tumor. For this purpose there are two interesting study, by Baumann *et al.* (30) and by Lindsay *et al.* (35) both targeting LM only from sarcoma. The most frequently prescribed doses were 50 Gy in 4–5 fractions for the first and 50 Gy in 10 fractions for the latter. The two-year LC was 86% for the first and 95% for the second.

In 2019, Hörner-Rieber *et al.* (36) published a multi-center study on a large number of oligo metastatic lung lesions (n=301) from non-small cell lung cancer (NSCLC) treated in 24 different German and Swiss centers between 1997 and 2017. The number and size of fractions were tailored to tumor size and location (peripheral *vs.* central). The authors report a BED range at isocenter of 128.2 Gy (37.5–323.4 Gy) for all PM with a difference between patients without an early progression 118.2 Gy (50.7–173.1 Gy) and with early progression 117.0 Gy (54.0–189.0 Gy). The primary value of the study is in the homogeneity of the sample in term of primary tumor, all NSCLC. At different time point of FU: 1-, 2-, 3-, and 5-year, LC was 91.9%, 82.0%, 76.4%, and 70.3%, respectively and OS was 80.8%, 62.2%, 48.1% and 28.8% respectively. This multi-institutional patterns-of-care analysis confirmed favorable LC and promising OS of SBRT for PM in patients with NSCLC.

A recent study on 316 metastases from various primary tumors by Kerstin A. Kessel *et al.* (39) confirms the promising role of SBRT in terms of LC and OS even with a long FU. They reported a low rate of severe side effects. These data confirm the strength of SBRT as therapeutic option and how it is competitive with more invasive treatment.

Summary

From our literature analysis, we can summarize:

- ❖ The ideal candidate for SBRT of lung metastases with curative intent is the oligometastatic patient according to the ESTRO/ASTRO definition.
- ❖ The patient management should be multidisciplinary,

in highly specialized cancer center, with the involvement of the different professional figures such as surgeons, medical oncologists, radiation oncologists, radiologists, pathologists and specialist in nuclear medicine.

- ❖ Patient's performance status should be preliminarily evaluated in order to select only those able to collaborate to a correct planning and delivery of the treatment.
- ❖ When planning the dose to be administered as effective and therefore “radical”, lesions should receive a minimum total BED of 100 Gy.
- ❖ The stereotactic technique, should allow the delivery the high dose to the target with a rapid dose fall off toward the OAR, particularly the intact lungs parenchyma.
- ❖ Each treatment fraction should be accurately monitored with all the equipment available to verify the precise implementation of what has been planned via the IGRT tools.
- ❖ All authors concludes that SBRT for pulmonary oligo metastases is safe and effective, with approximately 80% local control rates.
- ❖ Some hystotipes appear to respond less to the therapy such as metastases from colorectal, breast, pulmonary cancer and sarcomas. This assertion reflects a common feeling among the authors but is not supported by evidence from randomized studies on each primary tumor's histology.
- ❖ To date SBRT represents an alternative and competitive option for the cure of patients with oligo metastatic lung disease other than those unsuitable or refusing surgery.
- ❖ Future randomized studies will serve to better define the most appropriate dose and fractionation, the most suitable patients in terms of histology of the primary tumors and the best integration with systemic therapies.

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